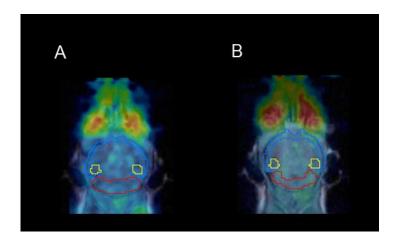
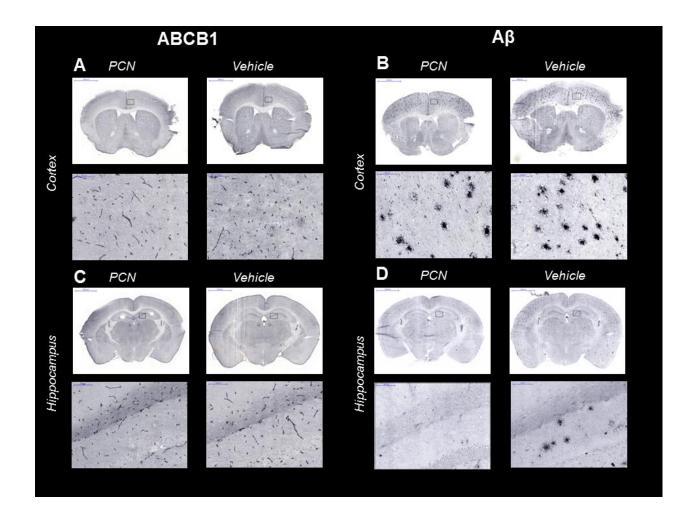


SUPPLEMENTAL FIGURE 1. Total radioactivity (mean±SD) in venous blood (A) and venous plasma (B) of samples collected at the end of the PET scan for wild-type and APP/PS1-21 mice aged 170 days at baseline and after treatment with PCN or vehicle. ns, not significant, one-way ANOVA followed by Tukey's multiple comparison test



SUPPLEMENTAL FIGURE 2. Representative coronal MR co-registered PET summation images (0-90 min) in one wild-type mouse (A) aged 170 days and one APP/PS1-21 mouse (B) aged 170 days. The outlined regions of interest are shown: cortex (blue), hippocampus (yellow) and cerebellum (red). The cerebellum served as a control region without Aβ deposits (but with ABCB1 expression).



SUPPLEMENTAL FIGURE 3. Immunohistochemical staining of ABCB1 in brain microvessels (A, C) and A β (B, D) in the cortex and hippocampus of APP/PS1-21 mice aged 170 days treated with PCN or vehicle (scale bar=2000 μ m). An enlarged section of the area is shown at a 20x magnification below each image (scale bar=100 μ m).

SUPPLEMENTAL TABLE 1.

Metabolism of ¹¹C-Metoclopramide*

Wild-type [†]	APP/PS1-21 [†]	PCN-treated ^{††}
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n	3	4	3
Plasma	39±8	24±2 [§]	24±4 [§]
Brain	92±4	82±7	86±7
Liver	78±3	61±3§	64±6 [§]
Kidney	66±4	63±5	58±3
Bile	56±9	33±10	45±21
Urine	53±8	40±12	39±20

^{*}Stated is the mean±SD percentage of unchanged ¹¹C-metoclopramide in different tissues and fluids determined with radio-thin-layer chromatography at 15 min after injection of ¹¹C-metoclopramide

[†]Animals were i.p. pretreated for 7 days once daily with PCN vehicle (5% dimethyl sulfoxide in safflower oil)

^{††}Animals (wild-type: n=1, APP/PS1-21: n=2) were i.p. pretreated for 7 days once daily with PCN (25 mg/kg)

 $^{{}^{\}S}P$ <0.05, for comparison with wild-type, one-way ANOVA followed by a Tukey's multiple comparison test

SUPPLEMENTAL TABLE 2

 $k_{\rm E, brain}$ Values in All Examined Animal Groups

Group	Region	$k_{\mathrm{E,brain}} (1/\mathrm{h})$
	Hippocampus	0.650 ± 0.074
Wild-type baseline	Cortex	0.665 ± 0.063
	Cerebellum	0.775±0.093
	Hippocampus	1.003 ± 0.157
Wild-type PCN	Cortex	0.958 ± 0.155
	Cerebellum	1.059±0.145
	Hippocampus	0.722 ± 0.055
Wild-type vehicle	Cortex	0.758 ± 0.059
	Cerebellum	0.934±0.152
	Hippocampus	0.704 ± 0.094
APP/PS1-21 baseline	Cortex	0.660 ± 0.070
	Cerebellum	0.812±0.109
	Hippocampus	0.938 ± 0.083
APP/PS1-21 PCN	Cortex	0.918 ± 0.089
	Cerebellum	1.023±0.073
	Hippocampus	0.733 ± 0.100
APP/PS1-21 vehicle	Cortex	0.711 ± 0.093
	Cerebellum	0.841±0.115
	Hippocampus	0.882 ± 0.065
Wild-type baseline (50d)	Cortex	0.743 ± 0.044
	Cerebellum	0.893 ± 0.027
	Hippocampus	0.817±0.108
APP/PS1-21 baseline (50d)	Cortex	0.739 ± 0.088
	Cerebellum	0.830 ± 0.097

Values are reported as mean±standard deviation.

 $k_{\rm E,brain}$ (1/h), elimination slope for radioactivity washout from the brain